

Figure S1. Representative flow cytometry plots show increased markers of inflammation in miR-146-/-mice during aging (~15 months of age). A. Splenic CD4+ T cells have an activated phenotype in aged miR-146a-/- mice. T cell-specific deletion of miR-155 reverses the inflammatory phenotype in miR-146a-/- T cells as evidenced by lower percentages of CD44+CD62L(low) subset. B. The frequency of PD1+CXCR5+ Tfh cells is higher in aged miR-146a-/- mice compared to wild type (WT). Loss of miR-155 in T cells reduced the Tfh levels back to WT levels. C. Germinal center B cells (GL7+FAS+ within the B220+IgD(int) population) are elevated in miR-146a-/- mice upon aging. Deletion of the T cell-expressed miR-155 mostly blunts this increase observed in miR-146a-/- mice.

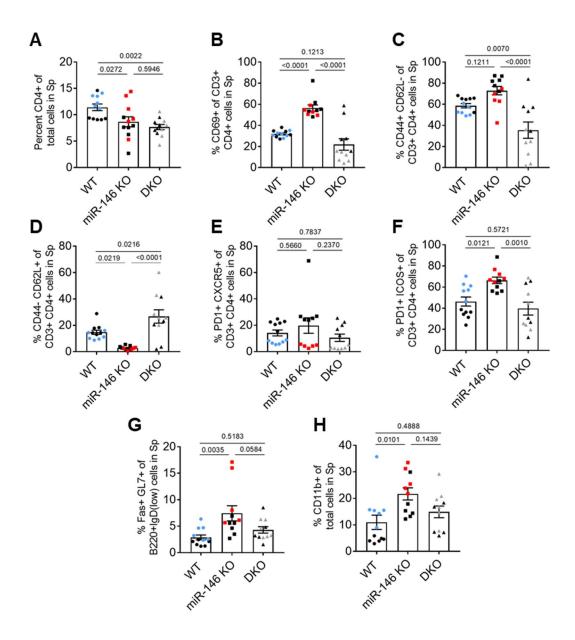


Figure S2. Flow cytometric assessment of the spleen reveals increased frequencies of inflammatory cells in aged miR-146a-/- mice (15 months of age). A. The frequency of CD4+ T cells within the total splenocytes (Sp) are shown. CD4+ T cells in aged miR-146a-/- mice exhibit an activated phenotype as evidenced by increased proportions of CD69+ (B) and CD44+CD62L- (C) subsets. D. The percentage of naïve T cells is reduced in miR-146a mice upon aging which is reversed back to wild type (WT) levels upon deletion of miR-155 in T cells. PD1+CXCR5+ (E) and PD1+ICOS+ (F) Tfh cells are elevated in aged miR-146a-/- mice. G. Germinal center B cell proportions increase in miR-146a-/- mice which was partially rescued by T cell-specific loss of miR-155. H. The frequency of CD11b+ myeloid cells is increased in aged miR-146a mice.

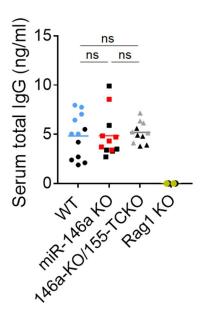
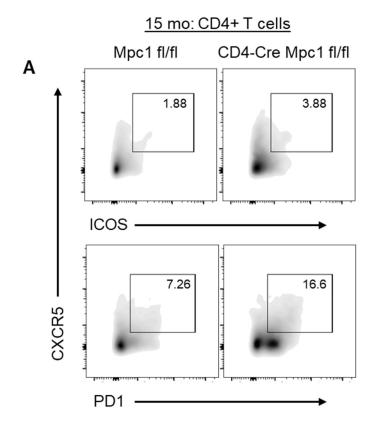


Figure S3. The overall IgG antibody levels in the serum are comparable between WT, miR-146a, and DKO mice. Unlike the levels of autoantibodies, total serum IgG levels from aged mice are not different among different groups, suggesting that aging is not associated with a non-specific increase in antibody production.



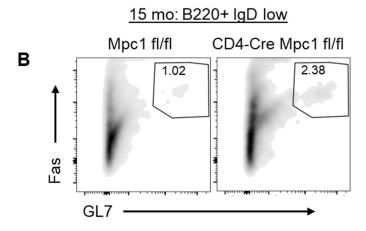


Figure S4. Representative flow cytometry plots showing splenic Tfh and GC B cells in aged Mpc1 T cell-conditional knockout mice (~15 months-old). A. The frequency of PD1+CXCR5+ and ICOS+CXCR5 Tfh cells is elevated upon T cell-specific deletion of Mpc1. B. Germinal center B cells are elevated in aged Mpc1 T cell-conditional knockout mice.